

SEQUENCE LISTING

<110> Warner-Lambert Company LLC
Bove, Susan R
Kilgore, Kenneth

<120> Methods of Treating Osteoarthritis with IL-6 Antagonists

<130> PC32145A

<150> 60/543,814

<151> 2004-02-11

<160> 12

<170> PatentIn version 3.3

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<211> 636

<212> DNA

<213> Mus musculus

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gaaatcgtgg aaatgagaaa agagttgtgc aatggcaatt ctgattgtat gaacaacgat 240
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gtccttcaga gagatacaga aactctaatt catatcttca accaagaggt aaaagattta 480
cataaaatag tccttcctac cccaatttcc aatgctctcc taacagataa gctggagtca 540
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<212> PRT

<213> Mus musculus

<400> 2

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35 40 45

Ser Gln Val Gly Gly Leu Ile Thr His Val Leu Trp Glu Ile Val Glu
50 55 60

Met Arg Lys Glu Leu Cys Asn Gly Asn Ser Asp Cys Met Asn Asn Asp
65 70 75 80

Asp Ala Leu Ala Glu Asn Asn Leu Lys Leu Pro Glu Ile Gln Arg Asn
85 90 95

Asp Gly Cys Tyr Gln Thr Gly Tyr Asn Gln Glu Ile Cys Leu Leu Lys
100 105 110

Ile Ser Ser Gly Leu Leu Glu Tyr His Ser Tyr Leu Glu Tyr Met Lys
115 120 125

Asn Asn Leu Lys Asp Asn Lys Lys Asp Lys Ala Arg Val Leu Gln Arg
130 135 140

Asp Thr Glu Thr Leu Ile His Ile Phe Asn Gln Glu Val Lys Asp Leu
145 150 155 160

His Lys Ile Val Leu Pro Thr Pro Ile Ser Asn Ala Leu Leu Thr Asp
165 170 175

Lys Leu Glu Ser Gln Lys Glu Trp Leu Arg Thr Lys Thr Ile Gln Phe
180 185 190

Ile Leu Lys Ser Leu Glu Glu Phe Leu Lys Val Thr Leu Arg Ser Thr
195 200 205

Arg Gln Thr
210

<210> 3
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<212> DNA
<213> Rattus norvegicus

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cacaacagac cagtatatac cacttcacaa gtcggaggct taattacata tgttctcagg 180
gagatcttgg aaatgagaaa agagttgtgc aatggcaatt ctgattgtat gaacagcgat 240
gatgcactgt cagaaaacaa tctgaaactt ccagaaatac aaagaaatga tggatgcttc 300
caaactggat ataaccagga aatttgcccta ttgaaaatct gctctggtct tctggagttc 360

cgtttctacc tggagtttgt gaagaacaac ttacaagata acaagaaaga caaagccaga 420
 gtcattcaga gcaatactga aaccctagtt catatcttca aacaagagat aaaagactca 480
 tataaaatag tccttcctac cccaacttcc aatgctctcc taatggagaa gtttagagtca 540
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 <213> Rattus norvegicus

<400> 4

Met Lys Phe Leu Ser Ala Arg Asp Phe Gln Pro Val Ala Phe Leu Gly
 1 5 10 15

Leu Met Leu Leu Thr Ala Thr Ala Phe Pro Thr Ser Gln Val Arg Arg
 20 25 30

Gly Asp Phe Thr Glu Asp Thr Thr His Asn Arg Pro Val Tyr Thr Thr
 35 40 45

Ser Gln Val Gly Gly Leu Ile Thr Tyr Val Leu Arg Glu Ile Leu Glu
 50 55 60

Met Arg Lys Glu Leu Cys Asn Gly Asn Ser Asp Cys Met Asn Ser Asp
 65 70 75 80

Asp Ala Leu Ser Glu Asn Asn Leu Lys Leu Pro Glu Ile Gln Arg Asn
 85 90 95

Asp Gly Cys Phe Gln Thr Gly Tyr Asn Gln Glu Ile Cys Leu Leu Lys
 100 105 110

Ile Cys Ser Gly Leu Leu Glu Phe Arg Phe Tyr Leu Glu Phe Val Lys
 115 120 125

Asn Asn Leu Gln Asp Asn Lys Lys Asp Lys Ala Arg Val Ile Gln Ser
 130 135 140

Asn Thr Glu Thr Leu Val His Ile Phe Lys Gln Glu Ile Lys Asp Ser
 145 150 155 160

Tyr Lys Ile Val Leu Pro Thr Pro Thr Ser Asn Ala Leu Leu Met Glu
 165 170 175

Lys Leu Glu Ser Gln Lys Glu Trp Leu Arg Thr Lys Thr Ile Gln Leu
 180 185 190

Ile Leu Lys Ala Leu Glu Glu Phe Leu Lys Val Thr Met Arg Ser Thr
 195 200 205

Arg Gln Thr
 210

<210> 5
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 <212> DNA
 <213> Human

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 <212> PRT
 <213> Human

<400> 6

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Gly Glu Asp Ser Lys Asp Val Ala Ala Pro His Arg Gln Pro Leu Thr
 35 40 45

Ser Ser Glu Arg Ile Asp Lys Gln Ile Arg Tyr Ile Leu Asp Gly Ile
 50 55 60

Ser Ala Leu Arg Lys Glu Thr Cys Asn Lys Ser Asn Met Cys Glu Ser
65 70 75 80

Ser Lys Glu Ala Leu Ala Glu Asn Asn Leu Asn Leu Pro Lys Met Ala
85 90 95

Glu Lys Asp Gly Cys Phe Gln Ser Gly Phe Asn Glu Glu Thr Cys Leu
100 105 110

Val Lys Ile Ile Thr Gly Leu Leu Glu Phe Glu Val Tyr Leu Glu Tyr
115 120 125

Leu Gln Asn Arg Phe Glu Ser Ser Glu Glu Gln Ala Arg Ala Val Gln
130 135 140

Met Ser Thr Lys Val Leu Ile Gln Phe Leu Gln Lys Lys Ala Lys Asn
145 150 155 160

Leu Asp Ala Ile Thr Thr Pro Asp Pro Thr Thr Asn Ala Ser Leu Leu
165 170 175

Thr Lys Leu Gln Ala Gln Asn Gln Trp Leu Gln Asp Met Thr Thr His
180 185 190

Leu Ile Leu Arg Ser Phe Lys Glu Phe Leu Gln Ser Ser Leu Arg Ala
195 200 205

Leu Arg Gln Met
210

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<211> 1323
<212> DNA
<213> Mus musculus

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gccaccgtta ccctgatttg ccccggaag gaagcagcag gcaatgttac cattcactgg 180
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gacgtgcagc tcagcgacac tggggactat ttatgctccc tgaatgatca cctgggtggg 300
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<210> 8
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 <212> PRT
 <213> Mus musculus

<400> 8

Met Leu Thr Val Gly Cys Thr Leu Leu Val Ala Leu Leu Ala Ala Pro
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Gly Thr Val Thr Ser Leu Pro Gly Ala Thr Val Thr Leu Ile Cys Pro
35 40 45

Gly Lys Glu Ala Ala Gly Asn Val Thr Ile His Trp Val Tyr Ser Gly
50 55 60

Ser Gln Asn Arg Glu Trp Thr Thr Thr Gly Asn Thr Leu Val Leu Arg
65 70 75 80

Asp Val Gln Leu Ser Asp Thr Gly Asp Tyr Leu Cys Ser Leu Asn Asp
85 90 95

His Leu Val Gly Thr Val Pro Leu Leu Val Asp Val Pro Pro Glu Glu

100	105	110
Pro Lys Leu Ser Cys Phe Arg Lys Asn Pro Leu Val Asn Ala Ile Cys 115 120 125		
Glu Trp Arg Pro Ser Ser Thr Pro Ser Pro Thr Thr Lys Ala Val Leu 130 135 140		
Phe Ala Lys Lys Ile Asn Thr Thr Asn Gly Lys Ser Asp Phe Gln Val 145 150 155 160		
Pro Cys Gln Tyr Ser Gln Gln Leu Lys Ser Phe Ser Cys Gln Val Glu 165 170 175		
Ile Leu Glu Gly Asp Lys Val Tyr His Ile Val Ser Leu Cys Val Ala 180 185 190		
Asn Ser Val Gly Ser Lys Ser Ser His Asn Glu Ala Phe His Ser Leu 195 200 205		
Lys Met Val Gln Pro Asp Pro Pro Ala Asn Leu Val Val Ser Ala Ile 210 215 220		
Pro Gly Arg Pro Arg Trp Leu Lys Val Ser Trp Gln His Pro Glu Thr 225 230 235 240		
Trp Asp Pro Ser Tyr Tyr Leu Leu Gln Phe Gln Leu Arg Tyr Arg Pro 245 250 255		
Val Trp Ser Lys Glu Phe Thr Val Leu Leu Leu Pro Val Ala Gln Tyr 260 265 270		
Gln Cys Val Ile His Asp Ala Leu Arg Gly Val Lys His Val Val Gln 275 280 285		
Val Arg Gly Lys Glu Glu Leu Asp Leu Gly Gln Trp Ser Glu Trp Ser 290 295 300		
Pro Glu Val Thr Gly Thr Pro Trp Ile Ala Glu Pro Arg Thr Thr Pro 305 310 315 320		
Ala Gly Ile Leu Trp Asn Pro Thr Gln Val Ser Val Glu Asp Ser Ala 325 330 335		
Asn His Glu Asp Gln Tyr Glu Ser Ser Thr Glu Ala Thr Ser Val Leu 340 345 350		

Ala Pro Val Gln Glu Ser Ser Ser Met Ser Leu Pro
 355 360

<210> 9
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 <212> DNA
 <213> Rattus norvegicus

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 cccagataa 1389

<210> 10
 <211> 364
 <212> PRT
 <213> Rattus norvegicus

<400> 10

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 20 25 30

Gly Thr Val Thr Ser Leu Pro Gly Ala Thr Val Thr Leu Ile Cys Pro
 35 40 45

Gly Lys Glu Ala Ala Gly Asn Ala Thr Ile His Trp Val Tyr Ser Gly
 50 55 60

Ser Gln Ser Arg Glu Trp Thr Thr Thr Gly Asn Thr Leu Val Leu Arg
 65 70 75 80

Ala Val Gln Val Asn Asp Thr Gly His Tyr Leu Cys Phe Leu Asp Asp
 85 90 95

His Leu Val Gly Thr Val Pro Leu Leu Val Asp Val Pro Pro Glu Glu
 100 105 110

Pro Lys Leu Ser Cys Phe Arg Lys Asn Pro Leu Val Asn Ala Phe Cys
 115 120 125

Glu Trp His Pro Ser Ser Thr Pro Ser Pro Thr Thr Lys Ala Val Met
 130 135 140

Phe Ala Lys Lys Ile Asn Thr Thr Asn Gly Lys Ser Asp Phe Gln Val
 145 150 155 160

Pro Cys Gln Tyr Ser Gln Gln Leu Lys Ser Phe Ser Cys Glu Val Glu
 165 170 175

Ile Leu Glu Gly Asp Lys Val Tyr His Ile Val Ser Leu Cys Val Ala
 180 185 190

Asn Ser Val Gly Ser Arg Ser Ser His Asn Val Val Phe Gln Ser Leu
 195 200 205

Lys Met Val Gln Pro Asp Pro Pro Ala Asn Leu Val Val Ser Ala Ile
 210 215 220

Pro Gly Ser Leu Val Gly Ser Lys Ser Val Gly Lys Thr Leu Ser Pro
 225 230 235 240

Gly Thr Gln Val Thr Thr Cys Cys Asn Ser Ser Phe Asp Thr Asp Leu
245 250 255

Tyr Gly Gln Arg Thr Phe Thr Val Trp Pro Leu Gln Val Ala Gln His
260 265 270

Gln Cys Val Ile His Asp Ala Leu Arg Gly Val Lys His Val Val Gln
275 280 285

Val Arg Gly Lys Glu Glu Phe Asp Ile Gly Gln Trp Ser Lys Trp Ser
290 295 300

Pro Glu Val Thr Gly Thr Pro Trp Leu Ala Glu Pro Arg Thr Thr Pro
305 310 315 320

Ala Gly Ile Pro Gly Asn Pro Thr Gln Val Ser Val Glu Asp Tyr Asp
325 330 335

Asn His Glu Asp Gln Tyr Gly Ser Ser Thr Glu Ala Thr Ser Val Leu
340 345 350

Ala Pro Val Gln Gly Ser Ser Pro Ile Pro Leu Pro
355 360

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<212> DNA
<213> Human

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aacatcacag tactgccgt ggccagaaac ccccgctggc tcagtgtcac ctggcaagac 720
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 aatacagact acttcttccc cagatag 1407

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 <211> 365
 <212> PRT
 <213> Human

<400> 12

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20 25 30

Gly Val Leu Thr Ser Leu Pro Gly Asp Ser Val Thr Leu Thr Cys Pro
35 40 45

Gly Val Glu Pro Glu Asp Asn Ala Thr Val His Trp Val Leu Arg Lys
50 55 60

Pro Ala Ala Gly Ser His Pro Ser Arg Trp Ala Gly Met Gly Arg Arg
65 70 75 80

Leu Leu Leu Arg Ser Val Gln Leu His Asp Ser Gly Asn Tyr Ser Cys
85 90 95

Tyr Arg Ala Gly Arg Pro Ala Gly Thr Val His Leu Leu Val Asp Val
100 105 110

Pro Pro Glu Glu Pro Gln Leu Ser Cys Phe Arg Lys Ser Pro Leu Ser
115 120 125

Asn Val Val Cys Glu Trp Gly Pro Arg Ser Thr Pro Ser Leu Thr Thr
 130 135 140
 Lys Ala Val Leu Leu Val Arg Lys Phe Gln Asn Ser Pro Ala Glu Asp
 145 150 155 160
 Phe Gln Glu Pro Cys Gln Tyr Ser Gln Glu Ser Gln Lys Phe Ser Cys
 165 170 175
 Gln Leu Ala Val Pro Glu Gly Asp Ser Ser Phe Tyr Ile Val Ser Met
 180 185 190
 Cys Val Ala Ser Ser Val Gly Ser Lys Phe Ser Lys Thr Gln Thr Phe
 195 200 205
 Gln Gly Cys Gly Ile Leu Gln Pro Asp Pro Pro Ala Asn Ile Thr Val
 210 215 220
 Thr Ala Val Ala Arg Asn Pro Arg Trp Leu Ser Val Thr Trp Gln Asp
 225 230 235 240
 Pro His Ser Trp Asn Ser Ser Phe Tyr Arg Leu Arg Phe Glu Leu Arg
 245 250 255
 Tyr Arg Ala Glu Arg Ser Lys Thr Phe Thr Thr Trp Met Val Lys Asp
 260 265 270
 Leu Gln His His Cys Val Ile His Asp Ala Trp Ser Gly Leu Arg His
 275 280 285
 Val Val Gln Leu Arg Ala Gln Glu Glu Phe Gly Gln Gly Glu Trp Ser
 290 295 300
 Glu Trp Ser Pro Glu Ala Met Gly Thr Pro Trp Thr Glu Ser Arg Ser
 305 310 315 320
 Pro Pro Ala Glu Asn Glu Val Ser Thr Pro Met Gln Ala Leu Thr Thr
 325 330 335
 Asn Lys Asp Asp Asp Asn Ile Leu Phe Arg Asp Ser Ala Asn Ala Thr
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 Ser Leu Pro Val Gln Asp Ser Ser Ser Val Pro Leu Pro
 355 360 365